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IN THE CLAIMS

Please replace all previous listings, and versions, of the claims with the following claims, where added text is indicated by underlining and deleted text is indicated by strikethrough:

- 1. (Original) A compound, comprising a non-protein-binding moiety (NPBM) and at least one protein-binding group (PBG).
- 2. (Original) The compound of claim 1, wherein the NPBM is a polyol, sugar, amino acid, or dendrimer moiety.
- 3. (Original) The compound of claim 1, wherein the NPBM is a polyol moiety; and said polyol moiety is a sorbitol or mannitol moiety.
- 4. (Original) The compound of claim 1, wherein the NPBM is a sugar moiety; and said sugar moiety is a glucose, sucrose, or trehalose moiety.
- 5. (Original) The compound of claim 1, wherein the NPBM is an amino acid moiety; and said amino acid moiety is an arginine betaine, proline, or ectoine moiety.
- 6. (Original) The compound of claim 1, wherein the NPBM is a dendrimer moiety; and said dendrimer moiety is based on benzene, pentaerythritol, P(CH₂OH)₃, or TRIS.
- 7. (Original) The compound of any of claims 1-6, wherein the PBG is a urea, guanidinium ion, detergent, amino acid, denaturant, surfactant, polysorbate, polaxamer, citrate, chaotrope, or acetate group.
- 8. (Original) The compound of any of claims 1-6, wherein the PBG is a guanidinium ion.
- 9. (Original) The compound of any of claims 1-6, wherein the PBG is sodium dodecyl sulfate.

$$\begin{array}{c} RO \\ \\ R' \end{array} \begin{array}{c} O \\ \\ N \\ \\ N \end{array} \begin{array}{c} \\ NH_2 \\ \\ \end{array}$$

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wherein:

R is an electron pair, H, alkyl, aryl, heteroaryl, aralkyl, heteroaralkyl, or an alkali metal;

R' is H, alkyl, aryl, heteroaryl, aralkyl, heteroaralkyl, or (R")₃N;

R" is an electron pair, H, alkyl, aryl, heteroaryl, aralkyl, or heteroaralkyl;

W is O, NH₂⁺(halogen), or S; and

n is 1, 2, or 4-100.

- 11. (Original) The compound of claim 10, wherein R is an electron pair.
- 12. (Original) The compound of claim 10, wherein R' is H.
- 13. (Original) The compound of claim 10, wherein R' is (R")₃N.
- 14. (Original) The compound of claim 10, wherein R' is H_3N^+ .
- 15. (Original) The compound of claim 10, wherein W is NH₂⁺Cl⁻.
- 16. (Original) The compound of claim 10, wherein n is 1.
- 17. (Original) The compound of claim 10, wherein n is 2.
- 18. (Original) The compound of claim 10, wherein n is 4.
- 19. (Original) The compound of claim 10, wherein n is 5.

- 20. (Original) The compound of claim 10, wherein n is 6.
- 21. (Original) The compound of claim 10, wherein R is an electron pair, R' is H_3N^+ , W is $NH_2^+Cl^-$, and n is 1.
- 22. (Original) The compound of claim 10, wherein R is an electron pair, R' is H_3N^+ , W is $NH_2^+Cl^-$, and n is 2.
- 23. (Original) The compound of claim 10, wherein R is an electron pair, R' is H_3N^+ , W is $NH_2^+Cl^-$, and n is 4.
- 24. (Original) The compound of claim 10, wherein R is an electron pair, R' is H_3N^+ , W is $NH_2^+Cl^-$, and n is 5.
- 25. (Original) The compound of claim 10, wherein R is an electron pair, R' is H_3N^+ , W is $NH_2^+Cl^-$, and n is 6.
- 26. (Original) The compound of claim 10, wherein R is an electron pair, R' is H_3N^+ , W is O, and n is 1.
- 27. (Original) The compound of claim 10, wherein R is an electron pair, R' is H_3N^+ , W is O, and n is 2.
- 28. (Original) The compound of claim 10, wherein R is an electron pair, R' is H_3N^+ , W is O, and n is 4.
- 29. (Original) The compound of claim 10, wherein R is an electron pair, R' is H_3N^+ , W is O, and n is 5.

30. (Original) The compound of claim 10, wherein R is an electron pair, R' is H_3N^+ , W is O, and n is 6.

- 31. (Original) The compound of claim 10, wherein R is an electron pair, R' is H, W is $NH_2^+Cl^-$, and n is 1.
- 32. (Original) The compound of claim 10, wherein R is an electron pair, R' is H, W is $NH_2^+Cl^-$, and n is 2.
- 33. (Original) The compound of claim 10, wherein R is an electron pair, R' is H^+ , W is $NH_2^+Cl^-$, and n is 4.
- 34. (Original) The compound of claim 10, wherein R is an electron pair, R' is H, W is $NH_2^+Cl^-$, and n is 5.
- 35. (Original) The compound of claim 10, wherein R is an electron pair, R' is H, W is $NH_2^+Cl^-$, and n is 6.
- 36. (Original) The compound of claim 10, wherein R is an electron pair, R' is H, W is O, and n is 1.
- 37. (Original) The compound of claim 10, wherein R is an electron pair, R' is H, W is O, and n is 2.
- 38. (Original) The compound of claim 10, wherein R is an electron pair, R' is H, W is O, and n is 4.
- 39. (Original) The compound of claim 10, wherein R is an electron pair, R' is H, W is O, and n is 5.

40. (Original) The compound of claim 10, wherein R is an electron pair, R' is H, W is O, and n is 6.

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41. (Original) A compound selected from the group consisting of:

wherein, independently for each occurrence,

R is an electron pair, H, alkyl, aryl, heteroaryl, aralkyl, heteroaralkyl, an alkali metal, or $\text{CH}_2\text{Y};$

R' is H, a sugar radical, or CH₂Y;

n is an integer from 1 to 100, inclusive;

a is 1, 2, or 3;

X is $C(CH_2Y)_3$; and

Y is a protein binding group,

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wherein at least one Y is present in all compounds.

42. (Original) The compound of claim 41, wherein Y is a guanidinium ion.

43. (Original) A polymer of formula II, III, IV, V, VI, VII, VIII, or IX:

$$\mathsf{R} = \left\{ \begin{array}{c} \mathsf{O} \\ \mathsf{R}' \end{array} \right\}_{\mathsf{D}} \left\{ \begin{array}{c} \mathsf{N} \\ \mathsf{N} \\ \mathsf{N} \\ \mathsf{N} \\ \mathsf{N} \\ \mathsf{P} \end{array} \right\}_{\mathsf{D}} \mathsf{R}$$

II

wherein, independently for each occurrence:

R is an electron pair, H, alkyl, aryl, heteroaryl, aralkyl, heteroaralkyl, or an alkali metal;

R' is H, alkyl, aryl, heteroaryl, aralkyl, heteroaralkyl, or (R")₃N;

R" is an electron pair, H, alkyl, aryl, heteroaryl, aralkyl, or heteroaralkyl;

W is O, NH₂⁺(halogen), or S;

n is 1, 2, or 4-100; and

p is an integer from 2 to 1000 inclusive;

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wherein, independently for each occurrence,

R is H, alkyl, aryl, heteroaryl, aralkyl, heteroaralkyl, or an alkali metal, or CH₂Y;

p is an integer from 2 to 1000 inclusive; and

Y is a PBG, wherein at least one Y is present;

wherein, independently for each occurrence:

R is H, alkyl, aryl, heteroaryl, aralkyl, heteroaralkyl, or an alkali metal, or CH₂Y; R' is H, alkyl, aryl, heteroaryl, aralkyl, heteroaralkyl, or (R'')₃N; R'' is an electron pair, H, alkyl, aryl, heteroaryl, aralkyl, or heteroaralkyl; p is an integer from 2 to 1000 inclusive; and

Y is a PBG, wherein at least one Y is present;

$$\begin{array}{c|c}
R & O & H \\
\hline
N^{+}(R)_{3} & Y & P
\end{array}$$

wherein, independently for each occurrence:

R is H, alkyl, aryl, heteroaryl, aralkyl, heteroaralkyl, or an alkali metal, or CH₂Y; n is an integer from 1 to 100 inclusive; p is an integer from 2 to 1000 inclusive; and Y is a PBG;

$$R = \begin{bmatrix} 0 & Y & \\ 0 & N & \\ N & \end{bmatrix}_{p}$$

$$VI$$

wherein, independently for each occurrence,

R is H, alkyl, aryl, heteroaryl, aralkyl, heteroaralkyl, an alkali metal, or CH₂Y; n is an integer from 1 to 100, inclusive;

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a is 1, 2, or 3;

Y is a PBG; and

p is an integer from 2 to 1000, inclusive;

$$R = \left(\begin{array}{c} O \\ Y \\ \end{array} \right) \left(\begin{array}{c} H \\ N \\ \end{array} \right) \left(\begin{array}{c} R \\ N \\ \end{array} \right) \left(\begin{array}{c} R \\ \end{array} \right) \left(\begin{array}{$$

VII

wherein, independently for each occurrence,

 $R\ is\ H,\ alkyl,\ aryl,\ heteroaryl,\ aralkyl,\ heteroaralkyl,\ an\ alkali\ metal,\ or\ CH_{2}Y;$

n is an integer from 1 to 6, inclusive;

Y is a PBG; and

p is an integer from 2 to 1000, inclusive; or

$$\mathsf{R} \overbrace{\bigg|}_{p}^{\mathsf{R}}$$

VIII

wherein, independently for each occurrence,

R is H, OH, alkyl, alkoxy, aryl, heteroaryl, aralkyl, heteroaralkyl, -O-alkali metal, CH₂Y, OCH₂Y, or has a structure selected from the following:

a is 1, 2, or 3;

X is $C(CH_2Y)_3$;

Y is a PBG, wherein at least one Y is present; and p is an integer from 2 to 1000, inclusive; or

IX

wherein, individually for each occurrence:

R is an electron pair, H, alkyl, aryl, heteroaryl, aralkyl, heteroaralkyl, or an alkali metal; R' is a sidechain of an alpha-amino acid, wherein at least one instance of R' is the sidechain of arginine;

X is O or NR; and

p is an integer from 2 to 1000, inclusive.

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44. (Original) A method of screening compounds or polymers for the property of inhibiting protein aggregation in solution, comprising:

- a) computing a set of parameters utilizing molecular modeling based on compounds or polymers known to have the property of inhibiting protein aggregation;
 - b) applying those parameters to other compounds or polymers; and
 - c) choosing the compounds or polymers that meet the criteria of those parameters.
- 45. (Original) A method of preparing a compound or polymers having the property of protein aggregation inhibition in solution, comprising:
- a) computing a set of parameters utilizing molecular modeling based on compounds or polymers known to have the property of inhibiting protein aggregation;
- b) designing a compound or polymer having the property of protein aggregation inhibition in solution based on those parameters; and
- c) synthesizing the compound or polymer having the property of protein aggregation inhibition in solution.
- 46. (Original) A method of classifying a compound or polymer as either inhibitory of protein aggregation in solution or not inhibitory of protein aggregation in solution, comprising:
- a) computing a set of parameters utilizing molecular modeling based on compounds or polymers known to have the property of inhibiting protein aggregation;
 - b) applying those parameters to a compound or polymer; and
- c) classifying the compound or polymer that meet the criteria of those parameters as inhibitory of protein aggregation in solution.
- 47. (Original) A method of determining the preferential binding coefficient, Γ_{XP} , of an additive in a protein solution, comprising:
- a) determining the phase space trajectories of the protein, solvent, and additive using molecular dynamics;
- b) calculating the distance, r, between the center of mass for both the solvent molecule and additive molecule to the protein's van der Waals surface;

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c) determining the minimum distance, r^* , at which no significant differences between the local $(r = r^*)$ and bulk density are observed;

- d) determining which molecules lie within the distance, r*, from the protein surface and classifying these molecules as the local domain;
- e) determining which molecules lie outside the distance, r*, from the protein surface and classifying these molecules as the bulk domain;
- f) determining the instantaneous preferential binding coefficient, $\Gamma_{XP}(t)$, using the following formula:

$$\Gamma_{XP}(t) = n^{II}_{X} - n^{I}_{X} (n^{II}_{W} / n^{I}_{W})$$

wherein:

 n_{X}^{II} = the number of additive molecules in the bulk domain;

 n_{X}^{I} = the number of additive molecules in the local domain;

 n^{II}_{W} = the number of solvent molecules in the bulk domain; and

 n_{W}^{I} = the number of solvent molecules in the local domain; and

g) calculating the preferential binding coefficient, Γ_{XP} , as the time average of each of the values in step f) using the following formula:

$$\Gamma_{XP} = \frac{1}{t} \int_{0}^{t} \Gamma_{XP}(t') dt'.$$

48. (Currently amended) A method of suppressing or preventing aggregation of a protein in solution, comprising the step of combining in a solution the compound or (i) a polymer of formula IX any of claims 1 to 43

$$\frac{\left[\begin{array}{ccc} O & R \\ I & N \\ R' & \end{array}\right]_p}{IX}$$

wherein, individually for each occurrence:

R is an electron pair, H, alkyl, aryl, heteroaryl, aralkyl, heteroaralkyl, or an alkali metal;

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R' is a sidechain of an alpha-amino acid, wherein at least one instance of R' is the sidechain of arginine;

X is O or NR; and p is an integer from 2 to 1000, inclusive; and (ii) a protein.

- 49. (Original) The method of claim 48, wherein the protein is a recombinant protein.
- 50. (Original) The method of claim 48, wherein the protein is a recombinant antibody.
- 51. (Original) The method of claim 48, wherein the protein is a recombinant human antibody.
- 52. (Original) The method of claim 48, wherein the protein is a recombinant human protein.
- 53. (Original) The method of claim 48, wherein the protein is recombinant human insulin, recombinant human erythropoietin or a recombinant human interferon.
- 54. (Original) The method of claim 48, wherein the solution is an aqueous solution.
- 55. (Original) The method of claim 48, wherein the protein is a recombinant protein; and the solution is an aqueous solution.
- 56. (Original) The method of claim 48, wherein the protein is a recombinant human protein; and the solution is an aqueous solution.
- 57. (Original) A method of decreasing the toxicological risk associated with administering a protein to a mammal in need thereof, comprising the steps of adding to a first solution of a protein a compound or polymer of any of claims 1 to 43 to give a second solution; and administering to a mammal in need thereof a therapeutic amount of said second solution.

- 58. (Original) The method of claim 57, wherein the protein is a recombinant protein.
- 59. (Original) The method of claim 57, wherein the protein is a recombinant antibody.
- 60. (Original) The method of claim 57, wherein the protein is a recombinant human antibody.
- 61. (Original) The method of claim 57, wherein the protein is a recombinant mammalian protein.
- 62. (Original) The method of claim 57, wherein the protein is a recombinant human protein.
- 63. (Original) The method of claim 57, wherein the protein is recombinant human insulin, recombinant human erythropoietin or a recombinant human interferon.
- 64. (Original) The method of claim 57, wherein the first solution and the second solution are aqueous solutions.
- 65. (Original) The method of claim 57, wherein the protein is a recombinant protein; and the first solution and the second solution are aqueous solutions.
- 66. (Original) The method of claim 57, wherein the protein is a recombinant human antibody; and the first solution and the second solution are aqueous solutions.
- 67. (Original) The method of claim 57, wherein the protein is a recombinant human protein; and the first solution and the second solution are aqueous solutions.

68. (Original) A method of facilitating native folding of a recombinant protein in solution, comprising the step of combining in a solution a compound or polymer of any of claims 1 to 43 and a recombinant protein.

- 69. (Original) The method of claim 68, wherein the recombinant protein is a recombinant antibody.
- 70. (Original) The method of claim 68, wherein the recombinant protein is a recombinant human antibody.
- 71. (Original) The method of claim 68, wherein the recombinant protein is a recombinant mammalian protein.
- 72. (Original) The method of claim 68, wherein the recombinant protein is a recombinant human protein.
- 73. (Original) The method of claim 68, wherein the recombinant protein is recombinant human insulin, recombinant human erythropoietin or a recombinant human interferon.
- 74. (Original) The method of claim 68, wherein the solution is an aqueous solution.
- 75. (Original) The method of claim 68, wherein the recombinant protein is a recombinant human antibody; and the solution is an aqueous solution.
- 76. (Original) The method of claim 68, wherein the recombinant protein is a recombinant human protein; and the solution is an aqueous solution.